


Substitute for form 1449/PTO  <b>INFORMATION DISCLOSURE STATEMENT BY APPLICANT</b>  <i>(Use as many sheets as necessary)</i>				<b>Complete If Known</b>	
				Application Number	10/728,665
				Filing Date	December 5, 2003
				First Named Inventor	Satyanarayana MEDICHERLA
				Art Unit	1614
				Examiner Name	A. Marschel
Sheet	1	of	3	Attorney Docket Number	219002032800

U.S. PATENT DOCUMENTS					
Examiner Initials*	Cite No. <sup>1</sup>	Document Number Number-Kind Code <sup>2</sup> (if known)	Publication Date MM-DD-YYYY	Name of Patentee or Applicant of Cited Document	Pages, Columns, Lines, Where Relevant Passages or Relevant Figures Appear
FOREIGN PATENT DOCUMENTS					
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NON PATENT LITERATURE DOCUMENTS			
Examiner Initials*	Cite No. <sup>1</sup>	Include name of the author (in CAPITAL LETTERS), title of the article (when appropriate), title of the item (book, magazine, journal, serial, symposium, catalog, etc.), date, page(s), volume-issue number(s), publisher, city and/or country where published.	T <sup>2</sup>
T.P.T. 	1.	ADAMS et al., "Pyrimidinylimidazole Inhibitors of CSBP/P38 Kinase Demonstrating Decreased Inhibition of Hepatic Cytochrome P450 Enzymes," Bioorg. Med. Chem. Lett. 8:3111-3116 (1998)	
	2.	BADGER et al., "Pharmacological Profile of SB 203580, a Selective Inhibitor of Cytokine Suppressive Binding Protein/p38 Kinase, in Animal Models of Arthritis, Bone Resorption, Endotoxin Shock and Immune Function," J. Pharmacol. Exp. Ther. 279:1453-1461 (1996)	
	3.	BALASA et al., "Islet-Specific Expression of IL-10 Promotes Diabetes in Nonobese Diabetic Mice Independent of Fas, Perforin, TNF Receptor-1, and TNF Receptor-2 Molecules," J. Immunology 165:2841-2849 (2000)	
	4.	BLAIR, "Regulation of Glucose Transport and Glycogen Synthesis in L6 Muscle Cells During Oxidation Stress," J. Biol. Sci. 17:36293-36299 (1999)	
	5.	BURYSEK et al., "The Serine Protease Plasmin Triggers Expression of MCP-1 and CD40 in Human Primary Monocytes via Activation of p38 MAPK and Janus Kinase (JAK)/STAT Signaling Pathways," J. Biol. Sci. 17:36293-36299 (1999)	
	6.	CASTEELS et al., "Prevention of Autoimmune Destruction of Syngeneic Islet Grafts in Spontaneously Diabetic Nonobese Diabetic Mice by a Combination of a Vitamin D <sub>3</sub> Analog and Cyclosporine," Transplantation 65:1225-1232 (1998)	
	7.	CHEN et al., "Monocyte Chemoattractant Protein-1 is Expressed in Pancreatic Islets from Prediabetic NOD Mice and in Interleukin-1 $\beta$ -Exposed Human and Rat Islet Cells," Diabetologia 44:325-332 (2001)	
	8.	COLLIS et al., "RPR203494 a Pyrimidine Analogue of the p38 Inhibitor RPR200765A with an Improved In Vitro Potency," Bioorg. Med. Chem. Lett. 11:693-696 (2001)	
	9.	CRUMP, "The Honeymoon Period in Non-Insulin-Dependent Diabetes Mellitus," J. Family Practice 25:78-82 (1987)	
	10.	DEBUSSCHE et al., "Course of Pancreatic $\beta$ Cell Destruction in Prediabetic NOD Mice: A Histomorphometric Evaluation," Diabete & Metabolisme (Paris) 20:282-290 (1994)	

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/T.P.T./	11.	DE LASZLO et al., "Pyrroles and Other Heterocycles as Inhibitors of p38 Kinase," Bioorg. Med. Chem. Lett. 8:2698 (1998)	
	12.	FIJEN et al., "Suppression of the Clinical and Cytokine Response to Endotoxin by RWJ-67657, a p38 Mitogen-Activated Protein-Kinase Inhibitor, in Healthy Human Volunteers," Clin. Exp. Immunol. 124:16-20 (2001)	
	13.	GALLAGHER et al., "Regulation of Stress-Induced Cytokine Production by Pyridinylimidazoles Inhibition of CSBP Kinase," Bioorg. Med. Chem. 5:49-64 (1997)	
	14.	GOEBELER et al., "The MKK6/p38 Stress Kinase Cascade Is Critical for Tumor Necrosis Factor- $\alpha$ -Induced Expression of Monocyte-Chemoattractant Protein-1 in Endothelial Cells," Blood 93:857-865 (1999)	
	15.	HALE et al., "Differential Expression and Activation of p38 Mitogen-Activated Protein Kinase $\alpha$ , $\beta$ , $\gamma$ , and $\delta$ in Inflammatory Cell Lineages," J. Immunol. 162:4246-4252 (1999)	
	16.	HANCOCK et al., "Suppression of Insulinitis in Non-Obese Diabetic (NOD) Mice by Oral Insulin Administration Is Associated with Selective Expression of Interleukin-4 and -10, Transforming Growth Factor- $\beta$ , and Prostaglandin-E," Am. J. Patho. 147:1194-1199 (1995)	
	17.	HEINZE and THON, "Honeymoon Period in Insulin-Dependent Diabetes Mellitus," Pediatrician 12:208-212 (1985)	
	18.	HERLAAR and BROWN, "p38 MAPK Signalling Cascades in Inflammatory Disease," Molec. Med. Today 5:439-447 (1999)	
	19.	HOSKER and TURNER, "Insulin Treatment of Newly-Presenting Ketotic Diabetic Patients into the Honeymoon Period," Lancet 18:633-635 (1982)	
	20.	JACKSON et al., "Pharmacological Effects of SB 220025, a Selective Inhibitor of P38 Mitogen-Activated Protein Kinase, in Angiogenesis and Chronic Inflammatory Disease Models," J. Pharmacol. Exp. Ther. 284:687-692 (1998)	
	21.	KEYMEULEN and SOMERS, "Immunointervention in Type 1 (Insulin-Dependent) Diabetes Mellitus," Acta Clinica Belgica 48:86-95 (1993)	
	22.	KIMBLE et al., "Simultaneous Block of Interleukin-1 and Tumor Necrosis Factor Is Required to Completely Prevent Bone Loss in the Early Postovariectomy Period," Endocrinol., 136:3054-3061 (1995)	
	23.	KUMAR et al., "Novel Homologues of CSBP/p38 MAP Kinase: Activation, Substrate Specificity and Sensitivity to Inhibition by Pyridinyl Imidazoles," Biochem. Biophys. Res. Comm. 235:533-538 (1997)	
	24.	LEE et al., "Inhibition of p38 MAP Kinase as a Therapeutic Strategy," Immunopharmacology 47:185-201 (2000)	
	25.	MAHON et al., "Lessons Learned from Use of Cyclosporine for Insulin-Dependent Diabetes Mellitus," Ann. N.Y. Acad. Sci. 696:351-363 (1993)	
	26.	MAKINO et al., "Breeding of a Non-Obese, Diabetic Strain of Mice," Exp. Animal 29:1-13 (1980)	
	27.	MCLAY et al., "The Discovery of RPR 200765A, p38 MAP Kinase Inhibitor Displaying a Good Oral Anti-Arthritic Efficacy," Bioorg. Med. Chem. 9:537-554 (2001)	
	28.	MIYAZAKI et al., "Predominance of T Lymphocytes in Pancreatic Islets and Spleen of Pre-Diabetic Non-Obese Diabetic (NOD) Mice: A Longitudinal Study," Clin. Exp. Immuno. 60:622-630 (1985)	
↓	29.	MORI et al., "Preventive Effects of Cyclosporin on Diabetes in NOD Mice," Diabetologia 29:244-247 (1986)	
	30.	PALMER and MCCULLOCH, "Perspective in Diabetes, Prediction and Prevention of IDDM-	

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<u>/T.P.T.</u>		1991," Diabetes 40:943-947 (1990)	
	31.	PAVLOVIC et al., "Activation of Extracellular Signal-Regulated Kinase (ERK) 1/2 Contributes to Cytokine-Induced Apoptosis in Purified Rat Pancreatic $\beta$ -Cells," Eur. Cytokine Netw. 11:267-274 (2000)	
	32.	POSTE et al., "Lipid Vesicles as Carriers for Introducing Biologically Active Materials into Cells," Meth. Cell Biol. 14:33 (1976)	
	33.	RAPOPORT et al., "Thymic T Cell Anergy in Autoimmune Nonobese Diabetic Mice Is Mediated by Deficient T Cell Receptor Regulation of the Pathway of p21 <sup>ras</sup> Activation," J. Exp. Med. 177:1221-1227 (1993)	
	34.	RAVELLI, "Macrophage Activation Syndrome," Curr. Opin. Rheumatol. 14:548-552 (2002)	
	35.	REVESZ et al., "SAR of 4-Hydroxypiperidine and Hydroxyalkyl Substituted Heterocycles as Novel p38 Map Kinase Inhibitors," Bioorg. Med. Chem. Lett., 10:1261-1264 (2000)	
	36.	ROGERS and GIEMBYCZ, "Conquering Airway Inflammation in the 21st Century," Drug Discov. Today 3:532-535 (1998)	
	37.	RYU et al., "Reversal of Established Autoimmune Diabetes by Restoration of Endogenous $\beta$ Cell Function," J. Clin. Invest. 108:63-72 (2001)	
	38.	SALOMON et al., "B7/CD28 Costimulation Is Essential for the Homeostasis of the CD4 <sup>+</sup> CD25 <sup>+</sup> Immunoregulatory T Cells that Control Autoimmune Diabetes," Immunity 12:431-437 (2000)	
	39.	SHAPIRO et al., "Combination Therapy with Low Dose Sirolimus and Tacrolimus is Synergistic in Preventing Spontaneous and Recurrent Autoimmune Diabetes in Non-Obese Diabetic Mice," Diabetologia 45:224-230 (2002)	
	40.	SPRINKEL et al., "Glucose Potentiates Interleukin-1 $\beta$ (IL-1 $\beta$ )-Induced p38 Mitogen-Activated Protein Kinase Activity in Rat Pancreatic Islets of Langerhans," Eur. Cytokine Netw. 12:331-339 (2001)	
	41.	SUGIHARA et al., "Ultrastructural and Immunoelectron Microscopic Studies on Infiltrating Mononuclear Cells in Lymphocytic Submandibulitis in NOD Mice," Histol. Histopathol. 4:397-404 (1989)	
	42.	TABATABAIE et al., "COX-2 Inhibition Prevents Insulin-Dependent Diabetes in Low-Dose Streptozotocin-Treated Mice," Biochem. Biophys. Res. Comm. 273:699-704 (2000)	
	43.	THOMAS and KAY, "Beta Cell Destruction in the Development of Autoimmune Diabetes in the Non-Obese Diabetic (NOD) Mice," Diabetes/Metabolism Res. Rev. 16:251-261 (2000)	
	44.	WADSWORTH et al., "RWJ 67657, a Potent, Orally Active Inhibitor of p38 Mitogen-Activated Protein Kinase," J. Pharmacol. Expt. Therapeut. 291:680-687 (1999)	
	45.	WANG et al., "Structural Basis of Inhibitor Selectivity in MAP Kinases," Structure 6:1117-1128 (1998)	
	46.	YOSHIDA and KIKUTANI, "Genetic and Immunological Basis of Autoimmune Diabetes in the NOD Mouse," Reviews in Immunogenetics 2:140-146 (2000)	
	47.	ZHANG et al., "CD28 Co-Stimulation Restores T Cell Responsiveness in NOD Mice by Overcoming Deficiencies in Rac-1/p38 Mitogen-Activated Protein Kinase Signaling and IL-2 and IL-4 Gene Transcription," International Immunology 13:377-384 (2001)	

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